700			U.S. Patent and Tra	PTO/SB/21 (02-04) pproved for use through 07/31/2006. OMB 0651-0031 demark Office: U.S. DEPARTMENT OF COMMERCE
Under the Paperwork Reduction Act of 1995, no person			Application Number	nformation unless it displays a valid OMB control number.  09/920,310
FORM  (to be used for all correspondence after initial filing)			Filing Date	August 1, 2001
			First Named Inventor	Nicolau
			Art Unit	1626
				· · · · · · · · · · · · · · · · · · ·
Total Number of Pages in This Submission 7			Examiner Name	Anderson, R.L.
Total Number of Pages III This Submission /			Attorney Docket Number	GMX-005.01
ENCLOSURES (check all that apply)				
Fee Transmittal Form		☐ Drawing(s)		After Allowance Communication to Group
Fee Attached		Licensing-related Papers		Appeal Communication to Board of Appeals and Interferences
Amendment / Reply		Petition		Appeal Communication to Group (Appeal Notice, Brief, Reply Brief)
After Final		Petition to Convert to a Provisional Application		Proprietary Information
Affidavits/declaration(s)		Power of Attorney, Revocation Change of Correspondence Address		Status Letter
Extension of Time Request		Terminal Disclaimer		Other Enclosure(s) (please identify below):
Express Abandonment Request		Request for Refund  CD, Number of CD(s)		Response to Non-Responsive Reply (6 pages); Return postcard
☐ Information Disclosure Statement		CD, Number of CD(s)		F
Certified Copy of Priority Document(s)		Remarks		
Response to Missing Parts/ Incomplete Application				
Response to Missing Parts under 37 CFR 1.52 or 1.53				
SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT				
Firm or Individual name	Michael DiVerdi			
Signature	Michael J. D. Verdi			
Date	June 17, 2004			
CERTIFICATE OF TRANSMISSION/MAILING				
Service with sufficie Alexandria, VA 22313	this correspondence is nt postage as first cl 3-1450 on the date sho	s being facs ass mail in	simile transmitted to the USPTO	or deposited with the United States Postal ommissioner for Patents, P.O. Box 1450,
Typed or printed name   Steve Pereira				

Signature

Date

June 17, 2004

This collection of information is required by 37 CFR 1.5. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Petent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

1626 Whu

In re Application of: Nicolau, Y.C. et al.

Serial No: 09/920,310

Filed: August 1, 2001

For: Ammonium Salts Of Hemoglobin Allosteric Effectors, and Uses Thereof

Attorney Docket No.: GMX-005.01

Examiner: Anderson, R. L.

Group Art Unit: 1626

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

## **Certificate of Mailing**

I hereby certify that this "Response to Non-Responsive Reply" is being deposited with the U. S. Postal Service as First Class Mail with sufficient postage on the date set forth below in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

6.17.09

Date of Signature and Mail Deposit

Steve Pereira

## Response to Non-Responsive Reply

Dear Examiner Anderson:

In a Non-Responsive Reply dated May 19, 2004, it was stated that the Applicants' selected species, made in a reply dated February 25, 2004, to a restriction requirement dated January 23, 2004, does not fall within elected Group I. The Applicants respectfully traverse this contention.

It was stated that the elected group requires an aliphatic ammonium cation and that the Applicants' election of a species having cyclohexyl ammonium cations does not satisfy this requirement because aliphatic by definition means an open chain hydrocarbon and not a cyclic one. The Applicants respectfully traverse this definition of aliphatic and respectfully submit that aliphatic includes both cyclic and open chain hydrocarbons.

To support their position, the Applicants have included pages 34 and 78 from Streitwieser and Heathcock's "Introduction to Organic Chemistry," third edition, 1985, which defines aliphatic as excluding aromatic compounds, but including both cyclic and open chain hydrocarbons.

Therefore, based on what one of ordinary skill in the art would consider aliphatic, as supported by text book definitions, the Applicants respectfully submit that their elected species comprising cyclohexyl ammonium cations falls within the elected group, and that the election made on February 25, 2004, was fully responsive.

#### <u>Fees</u>

The Applicants believe no fee is due in connection with the filing of this paper.

Nevertheless, the Director is hereby authorized to charge any required fee to our Deposit Account, 06-1448.

## Conclusion

The Applicants believe that they have responded fully to the Office communication dated May 19, 2004. However, if a telephone conversation with Applicant's Agent would expedite prosecution of the above-identified application, the Examiner is urged to contact the undersigned.

By: \_

Respectfully submitted, Foley Hoag LLP

Michael J. DiVerdi, PhD

Reg. No. 51,620

Agent for Applicants

155 Seaport Boulevard Boston, MA 02210

Telephone: (617) 832-1000 Telecopier: (617) 832-7000

Date: June 17, 2004

# INTRODUCTION to ORGANIC CHEMISTRY

Andrew Streitwieser, Jr. Clayton H. Heathcock

THIRD EDITION

# **BEST AVAILABLE CO**

THIS PAGE BLANK (USPTO)

AND COPY SO

Some Ethers

Chap. 3 Organic

Structures

$$CH_3$$
—O— $CH_3$   $CH_3$ —O— $CH_3$   $H_2$ C— $CH_2$   $H_2$ C  $CH_2$ 

The carbon-oxygen double bond, the carbonyl group, is found in aldehydes and ketones. When the carbonyl group is bonded to an OH group, it becomes a carboxy group. Compounds containing this functional group are called carboxylic acids.

Some Aldehydes

Some Carboxylic Acids

O O 
$$CH_2$$
 $\parallel$ 
 $\parallel$ 
 $HCOH$   $CH_3COH$   $\dot{H_2}C$ — $CHCOOH$ 

Table 3.1 lists a number of the important functional groups. The structures and names of these groups should be committed to memory. They form an essential part of the language of organic chemistry. In our subsequent studies we will develop the chemistry of the individual functional groups in terms of structural and electronic theory, nomenclature (names), physical properties, the preparation from other functional groups, and the characteristic reactions that produce other groups.

Interconversions of functional groups constitute a large proportion of organic chemistry. After the individual groups have been studied, the effect of one group on another can be considered, for the organic chemistry of compounds with more than one functional group is not simply the sum of the parts. Groups affect each other, sometimes in complex ways. One of the reasons for studying the theory of organic chemistry is that the mutual interactions of functional groups can be understood.

The aromatic ring in Table 3.1 is written with three carbon-carbon double bonds. Nevertheless, we shall see later (Chapters 20 and 22) that compounds containing this ring system differ substantially in their chemistry from the alkenes. Compounds containing this ring system are known collectively as aromatic compounds. Compounds with no aromatic ring are known as aliphatic compounds.

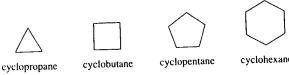
EXERCISE 3.2 Using R = ethyl, write structural and condensed formulas for one example each of an alkene, an alkyne, an alcohol, an ether, an aldehyde, a ketone, a carboxylic acid, an amine, and a nitrile.

HIS PAGE BLANKLESPTO

DEST AVAILABLE COPY

Chap. 5
Alkanes

Cycloalkanes are usually symbolized by simple geometric figures in which a carbon atom with its appropriate number of attached hydrogens is understood to be present at each apex. Thus, the four smallest cycloalkanes are depicted as



Simple substituted cycloalkanes are depicted by the appropriate geometric figure, with attached substituent groups.

1-chloro-3-methylcyclopentane

2-cyclohexyl-4-methylpentane

The alkanes and cycloalkanes are the parent structures in the general class of aliphatic compounds. Most of the chemistry of cycloalkanes is similar to that of the alkanes. There are some differences in stability and in their conformations, which will be discussed in Sections 5.6 and 7.7, respectively.

**EXERCISE 5.5** Using simple geometric figures and line structures, depict the following compounds. Compare your structures with complete structural representations.

(a) 1,1,3-trimethylcyclohexane

(b) 3-cyclopentylpentane

(c) 1-chloro-4-chloromethylcyclohexane

(d) 1,1,2,2-tetramethylcyclopropane

# 5.5 Heats of Formation

The heat of formation of a compound from its elements in their standard states is a thermodynamic property with considerable use in organic chemistry. This quantity, symbolized  $\Delta H_f^{\circ}$ , is defined as the enthalpy of the reaction of elements in their standard states to form the compound. The standard state of each element is generally the most stable state of that element at 25°C and 1 atm pressure. The standard state of carbon is taken as the graphite form, whereas those of hydrogen and oxygen are  $H_2$  and  $O_2$  gases, respectively. By definition,  $\Delta H_f^{\circ}$  for an element in its standard state is zero. The standard heat of formation of butane is  $-30.36 \pm 0.16$  kcal mole<sup>-1</sup> and that of 2-methylpropane is  $-32.41 \pm 0.13$  kcal mole<sup>-1</sup>.

4 C (graphite) + 5 
$$H_2(g) = n \cdot C_4 H_{10}(g)$$
  $\Delta H^{\circ} \equiv \Delta H_f^{\circ} (n \cdot C_4 H_{10})$   
= -30.36 kcal mole<sup>-1</sup>  
4 C (graphite) + 5  $H_2(g) = (CH_3)_3 CH(g)$   $\Delta H^{\circ} \equiv \Delta H_f^{\circ} (i \cdot C_4 H_{10})$   
= -32.41 kcal mole<sup>-1</sup>